

Supplementary Table 1

| Three-way ANOVA effect | F statistic | df | p-value |
|--|--------------------|--|------------------|
| CBD <i>versus</i> AM251 <i>versus</i> NH ₄ Cl interaction | 8.541 | (1,40) | <i>p</i> = 0.006 |
| CBD <i>versus</i> AM630 <i>versus</i> NH ₄ Cl interaction | 3.923 | (1,32) | <i>p</i> = 0.047 |
| CBD <i>versus</i> CPZ <i>versus</i> NH ₄ Cl interaction | 4.249 | (1,40) | <i>p</i> = 0.045 |
| Pairwise comparisons (Sidak's <i>post hoc</i> test) | | | |
| CBD versus AM251 versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM251 ⁽⁺⁾ / CTR | <i>p</i> = 0.024 |
| CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | <i>p</i> = 0.003 |
| CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | <i>p</i> = 0.001 |
| CBD versus AM630 versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM630 ⁽⁺⁾ / CTR | <i>p</i> = 0.018 |
| CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | <i>p</i> < 0.001 |
| CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | <i>p</i> < 0.001 |
| CBD versus CPZ versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / CPZ ⁽⁺⁾ / CTR | <i>p</i> < 0.001 |
| CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | <i>p</i> < 0.001 |
| CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | <i>p</i> < 0.001 |

Table S1. Statistical summary for CBD *versus* antagonists (AM251, AM630, CPZ) under the presence or not of NH₄Cl autophagic flux blockade in SH-SY5Y cells. Three-way ANOVA followed by Sidak's *post hoc* test for pairwise comparisons; respective *F* statistic, degree of freedom (*df*), and *p*-values are indicated in the table for every hypothesis test.

Supplementary Table 2

| Three-way ANOVA effect | F statistic | df | p-value |
|--|-------------|--|-----------|
| CBD versus AM251 versus NH ₄ Cl interaction | 4,352 | (1,32) | p = 0.045 |
| CBD versus AM630 versus NH ₄ Cl interaction | 6,674 | (1,32) | p = 0.015 |
| CBD versus CPZ versus NH ₄ Cl interaction | 5,498 | (1,32) | p = 0.025 |
| Pairwise comparisons (Sidak's post hoc test) | | | |
| CBD versus AM251 versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁻⁾ / AM251 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁻⁾ / CTR | p < 0.001 |
| CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM251 ⁽⁻⁾ / NH ₄ Cl | p = 0.001 |
| CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM251 ⁽⁺⁾ / NH ₄ Cl | p = 0.001 |
| CBD versus AM630 versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁻⁾ / AM630 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁻⁾ / CTR | p < 0.001 |
| CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / AM630 ⁽⁻⁾ / NH ₄ Cl | p < 0.001 |
| CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / AM630 ⁽⁺⁾ / NH ₄ Cl | p < 0.001 |
| CBD versus CPZ versus NH₄Cl experiment | | | |
| CBD 10 μM ⁽⁻⁾ / CPZ ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁻⁾ / CTR | p = 0.002 |
| CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁻⁾ / CPZ ⁽⁻⁾ / NH ₄ Cl | p = 0.001 |
| CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁻⁾ / NH ₄ Cl | vs | CBD 10 μM ⁽⁺⁾ / CPZ ⁽⁺⁾ / NH ₄ Cl | p = 0.001 |

Table S2. Statistical summary for CBD versus antagonists (AM251, AM630, CPZ) under the presence or not of NH₄Cl autophagic flux blockade in immortalized astrocytes cells. Three-way ANOVA followed by Sidak's *post hoc* test for pairwise comparisons; respective F statistic, degree of freedom (df), and p-values are indicated in the table for every hypothesis test.